

Aspects of the invention will now be further described with reference to the accompanying drawings in which:

Figures 1 to 6a/b are synthesis routes for compounds of formula **Ia** of the present invention;

Figures 7 to 14 are synthesis routes for compounds of formula **II** of the present invention;

Figures 15 to 24 are synthesis routes for compounds of formula **III** of the present invention;

Figure 25 is a synthesis route for a compound of formula **IV**;

Figure 26 is a synthesis of an intermediate in the preparation of compounds of formula **IV** of the present invention;

Figure 27 is a synthesis routes for compounds of formula **IV** of the present invention; and

Figures 28 to 31 are graphs illustrating the cytotoxicity results of examples 5 to 8 respectively.

Replace the title paragraph found on page 152, lines 14-16 with the following:

Example 3(g) : 8-Hydroxy-7,9-dimethoxy-1,2,3,11a-tetrahydropyrrolo[2,1-c][1,4]benzodiazepin-5-one (130, DRH-168)

Replace the title paragraph found on page 157, lines 22-23 with the following:

Examples 3(h) to (j) : Synthesis of 7-Phenyl PBDs (See Figure 21)

Replace the title paragraph found on page 164, lines 14-16 with the following:

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Example 3(k) : 8-Benzyloxy-7,9-dimethoxy-1,2,3,11a-tetrahydropyrrolo[2,1-c][1,4]benzodiazepin-5-one (143, DRH-105) (see Figure 22)

Replace the title paragraph found on page 166, lines 15-16 with the following:

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Example 3(l) : Synthesis of the C8-NH₂ PBD (151, AG/149) (see Figure 23)

Replace the title paragraph found on page 171, lines 4-6 with the following:

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Example 3(m) : Synthesis of (11aS)-8-methyl-7,9-dimethoxy-1,2,3,11a-tetrahydro-5H-pyrrolo[2,1-c][1,4]benzodiazepin-5-one (194) (see Figure 24)

Replace the title paragraph found on page 175, lines 11-15 with the following:

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Example 4 : Synthesis of the C8-Amines

Synthesis of 3-(11-Hydroxy-5-oxo-10-(2,2,2-trichloroethyloxocarbonylamino)-(11aS)-2,3,5,10,11,11a-hexahydro-1H-benzo[e]pyrrolo[2,1-a][1,4]diazepin-8-yloxy-2-propenylpropanoate (159) (see Figure 26)

Replace the title paragraph found on page 181, lines 1-3 with the following:

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Example 4(a) : 3-(7-methoxy-5-oxy(11aS)-2,3,5,11a-tetrahydro-1H-benzo[e]pyrrolo[1,2-a][1,4]diazepin-8-yloxy)-1-perhydro-1-pyrrolyl-1-propanone (161)(see Figure 27)

Replace the title paragraph found on page 183, lines 14-16 with the following:

Example 4(b) : 3-(7-methoxy-5-oxy(11aS)-2,3,5,11a-tetrahydro-1H-benzo[e]pyrrolo[1,2-a][1,4]diazepin-8-yloxy)-1-piperidino-1-propanone (163) (see Figure 27)

Replace the title paragraph found on page 185, lines 3-5 with the following:

Example 4(c) : 1,(2,3-dihydro-1H-indolyl)-3-(7-methoxy-5-oxy(11aS)-2,3,5,11a-tetrahydro-1H-benzo[e]pyrrolo[1,2-a][1,4]diazepin-8-yloxy)-1-propanone (165) (see Figure 27)

Replace the title paragraph found on page 187, lines 1-3 with the following:

Example 4(d) : 1,(2,3-dihydro-1H-2-isoindolyl)-3-(7-methoxy-5-oxy(11aS)-2,3,5,11a-tetrahydro-1H-benzo[e]pyrrolo[1,2-a][1,4]diazepin-8-yloxy)-1-propanone (167) (see Figure 27)

Replace the title paragraph found on page 189, lines 1-4 with the following:

Example 4(e) : Synthesis of (11aS) 8-(N-9-fluorenylmethoxycarbonyl)aminopropoxy-7-methoxy-1,2,3,11a-tetrahydro-5H-pyrrolo[2,1-c][1,4]benzodiazepin-5-one (205) (See Figure 25)

Replace the paragraph found on page 200, lines 28-31 with the following:

Example 5 : In Vitro Cytotoxicity of compounds of formula I

Some of the compounds synthesised in example 1, were subjected to the NCI *In Vitro* Cytotoxicity study. The results (LC₅₀;µM) are set out below, and are illustrated in Figure 28.